PCT/GB00/03228

CLAIMS

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- A method for producing a vaccine containing an immunogenic determinant, comprising the steps of:
- a) exposing extra-cellular pathogenic organisms to stress-inducing stimuli which would induce the production of SP/antigenic peptide fragment complexes;
 - b) extracting the endogenous stress-induced products from the treated organisms; and
 - c) using the extracted products as the immunogenic determinant in the preparation of the vaccine composition.
- 15 2. A method as claimed in claim 1, characterised in that the active ingredient of the immunogenic determinant consists predominantly of one or more shock protein/antigenic peptide fragment complexes.
- 20 3. A method as claimed in either of claims 1 or 2, characterised in that the stress-inducing stimulus is heat.
- 4. A method as claimed in claim 3, characterised in that the pathogenic organism is heated to from 5 to 8°C above the normal temperature for cultivation of the organisation.
- A method as claimed in any one of the preceding
 claims, characterised in that the pathogenic organism
 an extra-cellular procaryotic or protozoan

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species.

- 6. A method as claimed in any one of the preceding claims, characterised in that the pathogenic organism is a bacterial, protozoal or fungal species.
- 7. A method as claimed in any one of the preceding claims, characterised in immunogenic that the 10 determinant is а mixture of heat shock protein/antigenic peptide fragment complexes.
- 8. A method as claimed in any one of the preceding claims, characterised in that the extra-cellular pathogenic organism has been modified to induce or enhance the induction of the synthesis of stress proteins.
- 9. A method as claimed in any one of the preceding claims, characterised in that it is carried out in vitro.
 - 10.A method as claimed in claim 1, substantially as hereinbefore described in any one of the Examples.
 - 11.A vaccine composition containing an immunogenic determinant, characterised in that the immunogenic determinant comprises one or more complexes between a heat shock protein and an antigenic peptide fragment derived from the heat treatment of an extra-cellular pathogenic organisation.

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- 12.A vaccine composition produced by the method of any one of claims 1 to 10.
- 5 13.A vaccine composition as claimed in either of claims 11 or 12, characterised in that the composition also contains an adjuvant for the immunogenic determinant.
- 15.A vaccine composition as claimed in any one of claims
 11 to 14 substantially as hereinbefore described in any
 15 one of the Examples.
 - 16.A method for treating an animal with a vaccine, characterised in that it comprises administering a pharmaceutically acceptable quantity of a vaccine composition as claimed in any one of claims 11 to 15 sufficient to elicit an immune response in the animal.
- 17. A method for eliciting an immune response from an animal to infection by an intra-cellular pathogenic 25 organism which method comprises administering vaccine containing an immunogenic determinant. characterised in that the immunogenic determinant is an SP/antigenic peptide fragment complex produced in situ from the intra-cellular pathogen whose synthesis 30 is induced by external stress stimuli or by genetic

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modification of the pathogen so as to render its synthesis constitutive.